

Walter Jakob Gehring (1939-2014) [1]

By: Brinkman, Joe Keywords: [homeotic mutations](#) [2]

Walter Jakob Gehring [3] discovered the [homeobox](#) [4], a DNA segment found in a specific cluster of [genes](#) [5] that determine the body plan of animals, plants, and fungi. Gehring identified the [homeobox](#) [4] in 1983, with the help of colleagues while isolating the *Antennapedia* (*Antp*) gene in fruit flies (*Drosophila* [6]) at the [University of Basel](#) [7] in Basel, Switzerland. [Hox genes](#) [8], a family of [genes](#) [5] that contain the [homeobox](#) [4] sequence, determine the head-to-tail (anterior-posterior) body axis of both vertebrates and invertebrates. Gehring also identified the [homeobox](#) [4]-containing *Pax-6* gene as the master control gene in eye development of *Drosophila* [9], the same gene that, when mutated or absent in [humans](#) [10], leads to aniridia, or lack of the iris, in [humans](#) [10]. Gehring's work with the [homeobox](#) [4] suggested to biologists that widely different species share a similar and evolutionarily conserved genetic pathway that controls the development of overall body plans, from fruit flies to [humans](#) [10].

Gehring was born on 20 March 1939 in Zurich, Switzerland, to Marcelle Rembann and Jakob Gehring. Jakob Gehring was the chief engineer in turbine construction for the oceanliner Normandie. Gehring was the second child in the family and was younger than his sister by three years. Gehring later said that he and his sister got along very well and frequently competed against one another while growing up. When Gehring was a boy, his uncle sent him a cardboard box filled with pupae. The following spring, butterflies emerged from the pupae. Gehring stated that the event shaped his interest in animals and their development. Throughout his elementary school years, Gehring trapped [insects](#) [11] in the woods and brought them home to observe them.

Gehring studied [birds](#) [12] during his teenage years at Realgymnasium Zurich, a secondary school in Zurich, Switzerland. He studied the inheritance of the behaviors of migration and flight orientation, and he observed [birds](#) [12] by using the radar at the airport in Zurich. He attended the [University of Zurich](#) [13] in Zurich during the late 1950s and early 1960s, as an undergraduate zoology student, where developmental biologist [Ernst Hadorn](#) [14] mentored him. Gehring worked as a research assistant in Hadorn's lab, primarily focused on what mechanisms governed *Drosophila* [9] development. Hadorn allowed Gehring to use his radar data on bird migration for Gehring's thesis. Gehring noted that [birds](#) [12] lacked orientation when they could no longer recognize the location of the sun as a result of overcast clouds. A lack of genetic tools and funding prevented Gehring from further investigations of bird migration and orientation, and he switched his focus from [birds](#) [12] to *Drosophila* [9].

Hadorn's work on early developmental genetic control of *Drosophila* [9] shaped Gehring's doctoral focus in 1964. While Gehring was searching for a thesis topic, a fellow graduate student and Hadorn discovered a phenomenon, called transdetermination, in which embryonic cells switch from one developmental fate to another. An example that they found was cells fated to become antennae switched and became legs instead. Gehring soon after observed a genetic mutation, called Nasobemia, which transformed *Drosophila* [9] head antennae into a pair of legs. Gehring demonstrated that the gene neighbored *Antennapedia*, a gene necessary for leg development. Gehring then published a paper in which he argued that *Antennapedia* was responsible for activating the entire genetic pathway of leg development. During that same time in 1964, he married Elisabeth Lott. Their first son, Stephen, was born in 1965. Gehring and Lott had two children, with their second son Thomas born in 1970.

After earning his PhD in 1965, Gehring joined geneticist Alan Garen's lab as a postdoctoral fellow at [Yale University](#) [15] in New Haven, Connecticut, where he studied the molecular basis of structure development (homeotic) gene function. Gehring and Garen investigated the specification of cell fate, called [determination](#) [16], of cells in *Drosophila* [9] embryos. They noted that the cells could be determined at the [blastoderm](#) [17] stage, an early stage of development in which the embryonic sac is covered by the [blastoderm](#) [17], an outer layer of cells. The two used molecular markers to visualize a genetic fate map of the [blastoderm](#) [17] to see how cells developed structures in *Drosophila* [9]. They found that despite the [blastoderm](#) [17]'s homogeneous appearance, the cells within the [blastoderm](#) [17] were already determined for their respective cell fate, or lineage. As a result of his postdoctoral work, Gehring was appointed as an associate professor at the [Yale Medical School](#) [18] in 1969. He returned to Switzerland in 1972 and took a position as a professor in the department of cell biology at the Biozentrum of the [University of Basel](#) [7].

Gehring continued research with *Drosophila* [9] at the [University of Basel](#) [7] and focused on isolating the *Antennapedia* gene from *Drosophila* [9] during the 1970s. David Hogness, at [Stanford University](#) [19] in California, and his collaborators, developed a method scientists describe as walking along the chromosome, in which they clone any gene whose position on the chromosome they know. That method was used to successfully isolate the gene *Antennapedia*, a process that took three and a half years to complete. Once isolated, Gehring demonstrated that the gene *Antennapedia* shared common DNA sequences with its neighboring gene, *fushi tarazu*, which scientists had shown contributed to body [segmentation](#) [20] in the embryo.

In 1983, William McGinnis of Gehring's lab confirmed the homology, shared genetic sequences, between the [genes](#) [5] *Antennapedia* and *fushi tarazu*. He concluded that the homology was not spread across the entire gene, but only a 180 base pair segment. Gehring named the segment the [homeobox](#) [4], and Gehring used it as a probe to isolate entire sets of homeotic

proteins. By doing so, he provided evidence that homeotic [genes](#)^[5] encode for sequence-specific binding proteins and serve as control [genes](#)^[5] for body plan specification. Gehring and collaborators soon after observed that similar to [Drosophila](#)^[9], [mouse](#)^[21] [homeobox](#)^[4] [genes](#)^[5] were also clustered close together. Further evidence suggested comparable body plan pathways, due to the conservation of [homeobox](#)^[4] [genes](#)^[5], in both vertebrates and invertebrates, including [humans](#)^[10]. Some scientists claimed that the [homeobox](#)^[4] [genes](#)^[5], and their analogous role among species, provided a universal principle in the concept of development because [homeobox](#)^[4] [genes](#)^[5] determine body plan across broadly different species.

Further research conducted in Gehring's lab resulted in supplementary evidence regarding conserved [evolution](#)^[22] between species. One of Gehring's graduate students, Rebecca Quiring, accidentally cloned a [Drosophila](#)^[9] gene during a control experiment in 1994. The gene was homologous to the [mouse](#)^[21] *Pax-6* gene, which when mutated, corresponds to small eye mutations in mice. Mutations to that gene also causes aniridia in [humans](#)^[10], which is the lack of the iris, the structure that controls pupil dilation in the eye. Scientists observed that fly embryos that had the accidentally cloned gene, a [Drosophila](#)^[9] homolog of *Pax-6*, grew into adult flies that had no eyes. Gehring's collaborators then showed that the *Pax-6* gene could cause eyes to develop when expressed in other regions of [Drosophila](#)^[9]'s [genome](#)^[23]. The team expressed the *Pax-6* gene, and induced completely functioning eyes, on [Drosophila](#)^[9] legs, wings, and antennae. Gehring later claimed that those results indicated that eyes, from organisms in species as different as flies and mice, evolved from a common ancestor (monophyletic trait). The researchers claimed that the *Pax-6* developmental pathway appears to have been highly conserved evolutionarily, playing an integral role in the development of the prototypical eye.

Gehring became a professor emeritus in 2009 at the [University of Basel](#)^[7]. After his discovery of the [homeobox](#)^[4], research of its functionality continued in fungi, plants, and animals. Gehring continued to extrapolate [Drosophila](#)^[9] research to [humans](#)^[10], specifically for medicinal purposes. He died on 29 May 2014 as a result of an automobile accident.

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Walter Jakob Gehring discovered the homeobox, a DNA segment found in a specific cluster of genes that determine the body plan of animals, plants, and fungi. Gehring identified the homeobox in 1983, with the help of colleagues while isolating the Antennapedia (*Antp*) gene in fruit flies (*Drosophila*) at the University of Basel in Basel, Switzerland. Hox genes, a family of genes that have the homeobox, determine the head-to-tail (anterior-posterior) body axis of both vertebrates and invertebrates. Gehring also identified the homeobox-containing *Pax-6* gene as the master control gene in eye development of *Drosophila*, the same gene that, when mutated or absent in humans, leads to aniridia, or lack of the iris, in humans. Gehring's work with the homeobox suggested to biologists that widely different species share a similar and evolutionarily conserved genetic pathway that controls the development of overall body plans, from fruit flies to humans.

Subject

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